

# The SchizophreniaGene Database: A comprehensive online field synopsis and meta-analysis resource

Nicole C. Allen, Sachin Bagade, Matthew B. McQueen,  
John P. Ioannidis, Fotini K. Kavvoura, Muin J. Khouri, Rudolph E. Tanzi,  
Lars Bertram



**Genetics and Aging Research Unit**  
MassGeneral Institute for Neurodegenerative Diseases  
Massachusetts General Hospital  
Harvard Medical School



# SZ genetic association findings to date

- ⇒ >1,300 individual publications in scientific journals
- ⇒ >660 different genes have been tested (>5,000 variants)
- ⇒ ~20-30% show “significant association” in at least one study
- ⇒ Majority of positive associations never replicated (even once)
- ⇒ Currently up to 10 association papers published *each month*
- ⇒ Two published GWA studies (several more expected in 2008)

**Total number of *established* associations = 0**

Bertram L, McQueen MB, Mullin K, Blacker D, Tanzi RE.  
Systematic meta-analyses of Alzheimer disease genetic  
association studies: the AlzGene database. *Nat Genet*. 2007  
Jan;39(1):17-23.

# SzGene data collection and analysis

Initial search for papers June-July 2006  
on PubMed: “schizophreni\* & associat\*”  
**(now daily searches for “schizophreni”)**



~12,000 articles  
(currently: 300 per month)

Include if:

- a) genetic association study
- b) peer-reviewed (i.e. no abstracts)
- c) published in English

Enter study details into internal database  
and double-check all entries



>1,300 articles

Perform random-effects allelic  
meta-analyses if genotype data  
available from at least 4 case-  
control samples

Publish data online at:  
**www.szgene.org**



Current stats:  
1,281 studies, 661 loci  
138 meta-analyses

# SzGene literature search strategies are comprehensive (albeit not perfect)

Gene	SzGene	HuGENet	GAD	EMBASE
ACE	5	4*	3	2**
AKT1	9	6*	3	9
ASCL1	2	1	0	2
ATXN8OS	4	2	2	1
FZD3	7	5	4	7
IL1A	3	1	0	1
MLC1	6	2	2	2
PHOX2B	2	0	0	2
PLA2G4C	4	1	1	4
SOD2	5	5*	1	3
<b>Total</b>	<b>47</b>	<b>27</b>	<b>16</b>	<b>33</b>

\*Additional studies are listed on HuGENet that were not eligible for inclusion

\*\*One study found in EMBASE was not listed on PubMed

Note that HuGENet does not include studies published before 10/14/2000  
which applied to 3 out of the 47 studies included in SzGene

**WHAT'S NEW**

Recent Updates

**SRF PAPERS**

Current Papers

Search All Papers

Search Comments

**NEWS**

Research News

**FORUMS**

Current Hypotheses

Idea Lab

Online Discussions

Interviews

**RESOURCES**

What We Know

SchizophreniaGene

Drugs in Trials

Research Tools

Jobs

Conferences

Journals

General Information

**SRF COMMUNITY**

Member Directory

Researcher Profiles

Institutes and Labs

**ABOUT THE SITE**

Mission

History

SRF Team

Advisory Board

Support Us

How to Cite

The Schizophrenia Research Forum web site is sponsored by [NARSAD](#), the Mental Health Research Association, and supported in part by a contract from the [National Institute of Mental Health](#), National Institutes of Health, Department of Health and Human Services

**SchizophreniaGene (SZGene)**  
Published Candidate Genes for Schizophrenia

[BACK](#) [SEARCH](#) [METHODS](#) [DISCLAIMER](#) [CREDITS](#)



Updated 15 January 2008

Chromosome: [1](#) [2](#) [3](#) [4](#) [5](#) [6](#) [7](#) [8](#) [9](#) [10](#) [11](#) [12](#) [13](#)  
[14](#) [15](#) [16](#) [17](#) [18](#) [19](#) [20](#) [21](#) [22](#) [X](#) [Y](#)

Gene:  -- Select --

Protein:  -- Select --

Polymorphism:  -- Select --

Study:  -- Select --

Keyword:

\* New \* [View large scale studies \(including GWA analyses\)](#)

[Display, print, and download the SchizophreniaGene database information](#)

The SchizophreniaGene database aims to provide a comprehensive unbiased and regularly updated collection of genetic association studies performed on schizophrenia phenotypes. Eligible publications are identified following systematic searches of scientific literature databases as well as the table of contents of journals in genetics and psychiatry.

The database can be searched either by a variety of dropdown menus or by specific keywords. For each gene, summary overviews are provided displaying key characteristics for each publication, including links to genotype distributions of the polymorphisms studied, random-effect allelic meta-analyses, and funnel plots for an assessment of publication bias.

For more details on the background and methods, see [Methods](#), [Disclaimer](#), and [Credits](#). We encourage authors and readers to [contact us](#) to report errors in the presentation of study details, or to notify us of studies that have mistakenly been left out.

**How to Cite Content on SchizophreniaGene:**

Allen NC, Bagade S, Tanzi R, Bertram L. The SchizophreniaGene Database. Schizophrenia Research Forum. Available at: <http://www.schizophreniaforum.org/res/sczgene/default.asp>. Accessed [date of access].

**SchizophreniaGene**  
Top Results

[View Top Results Methods](#)

1. [DRD2](#)
2. [GRIN2B](#)
3. [GABRB2](#)
4. [PLXNA2](#)
5. [DTNBP1](#)
6. [TPH1](#)
7. [DRD4](#)
8. [IL1B](#)
9. [DAOA](#)
10. [DRD1](#) close
11. [APOE](#)
12. [MTHFR](#)
13. [SLC6A4](#)
14. [HP](#)
15. [DAO](#)
16. [TP53](#)
17. [AKT1](#)
18. [COMT](#)
19. [PPP3CC](#)

# **Summary of SzGene findings (April 30<sup>th</sup> 2007)**

- ⇒ Total number of papers = 1,179 (reporting on 516 genes)
- ⇒ Meta-analyses on 118 polymorphisms (52 genes)
- ⇒ Median sample size = 3,589 (IQR: 2,335-5,669)
- ⇒ Median number of studies = 6 (IQR: 4-9)

## **“Positive” associations (nominal P≤0.05)**

- ⇒ 24 variants in 16 genes
- ⇒ Average OR = 1.23 (range: 1.11-1.52; P 0.048 to <0.0001)
- ⇒ Median sample size = 3,378 (IQR: 2,410-5,419)

## **“Negative” associations (nominal P>0.05)**

- ⇒ 94 variants in 45 genes
- ⇒ Median sample size = 3,928 (IQR: 2,335-56,695)

# Significant findings in SzGene (April 2007)

Gene	Polymorphism	Model	Cases vs. controls (# independent samples)	OR (95% CI)†	P-value	Grade††
APOE	APOE ( $\epsilon$ 2/3/4) E4 vs. E3	E4 vs. E3, Caucasian <sup>a</sup>	1500 vs. 2702 (15)	1.16 (1.00-1.34)	0.043	B
COMT	rs165599	G vs. A, all ethnicities	2628 vs. 7340 (6)	1.11 (1.02-1.21)	0.019	C
COMT	rs737865	C vs. T, Caucasian <sup>a</sup>	1605 vs. 4021 (3)	1.13 (1.01-1.28)	0.039	C
DAO	rs4623951	C vs. T, all ethnicities	1509 vs. 1521 (4)	0.88 (0.79-0.98)	0.026	C
DRD1	rs4532 (DRD1_48A/G)	G vs. A, all ethnicities	725 vs. 1075 (5)	1.18 (1.01-1.38)	0.037	A
DRD2	rs1801028 (Ser311Cys)	G vs. C, Caucasian <sup>b</sup>	2299 vs. 3777 (15)	1.52 (1.09-2.12)	0.013	B
DRD2	rs6277 (Pro319Pro)	C vs. T, Caucasian <sup>b</sup>	473 vs. 896 (3)	1.45 (1.21-1.73)	<0.00004	C
DRD4	rs1800955 (521T/C)	C vs. T, all ethnicities	2002 vs. 1986 (6)	1.15 (1.05-1.26)	0.003	C
DRD4	120-bp TR	S vs. L, all ethnicities	1236 vs. 1199 (4)	0.81 (0.70-0.94)	0.005	C
DTNBP1	rs1011313 (P1325)	T vs. C, Caucasian <sup>a</sup>	2696 vs. 2849 (8)	1.23 (1.07-1.40)	0.003	A
GABRB2	rs1816072	C vs. T, Caucasian <sup>a</sup>	1129 vs. 995 (4)	0.82 (0.72-0.93)	0.002	C
GABRB2	rs1816071	G vs. A, Caucasian <sup>a</sup>	1133 vs. 993 (4)	0.82 (0.72-0.93)	0.002	C
GABRB2	rs194072	C vs. T, Caucasian <sup>a</sup>	1137 vs. 991 (4)	0.83 (0.69-1.00)	0.048	B
GABRB2	rs6556547	T vs. G, Caucasian <sup>a</sup>	774 vs. 620 (3)	0.70 (0.52-0.95)	0.022	B
GRIN2B	rs7301328 (366G/C)	G vs. C, all ethnicities	903 vs. 810 (4)	1.16 (1.01-1.33)	0.034	C
GRIN2B	rs1019385 (200T/G)	G vs. T, all ethnicities	502 vs. 466 (4)	1.45 (1.14-1.85)	0.003	C
HP	Hpl2	1 vs. 2, all ethnicities	1346 vs. 2018 (6)	0.88 (0.80-0.98)	0.016	C
IL1B	rs16944 (C511T)	T vs. C, Caucasian <sup>b</sup>	819 vs. 1302 (5)	0.78 (0.65-0.93)	0.006	C
MTHFR	rs1801133 (C677T)	T vs. C, all ethnicities	3327 vs. 4093 (14)	1.16 (1.05-1.30)	0.005	C
MTHFR	rs1801131 (A1298C)	C vs. A, Caucasian <sup>b</sup>	1211 vs. 1729 (5)	1.19 (1.07-1.34)	0.002	A
PLXNA2	rs752016	C vs. T, all ethnicities	1122 vs. 1211 (6)	0.82 (0.69-0.99)	0.037	C
SLC6A4	5-HTVNTR	10 vs. 12, all ethnicities	2335 vs. 2688 (11)	0.86 (0.74-0.99)	0.036	C
TP53	rs1042522	C vs. G, all ethnicities	1418 vs. 1410 (5)	1.13 (1.01-1.26)	0.029	C
TPH1	rs1800532 (218A/C)	A vs. C, all ethnicities	829 vs. 1268 (5)	1.31 (1.15-1.51)	<0.00008	A

# **Grading of epidemiologic credibility (HuGENet Venice interim guidelines)**

## **1. “Amount of evidence”**

⇒ sample size (>1000, 1000-100, <100)

## **2. “Replication”**

⇒ degree of heterogeneity ( $I^2$ -statistic)

## **3. “Protection from bias”**

⇒ OR <1.15, publication bias, sensitivity (initial, HWE)

***Lowest score in any category determines overall grade***

# Applying the HuGENet guidelines to SzGene

**Supplementary Table 6:** Details of the application of grading scheme for the assessment of cumulative evidence in genetic association studies (Venice interim criteria) on all SNPs showing nominally significant results in meta-analyses of all ethnicities or Caucasian-only

Shaded columns denote criteria which make up "OVERALL" score (see main text for details)

"A" = strong, "B" = moderate, "C" = weak.

Gene	Ethnicity	Polymorphism	AMOUNT	N minor	REPLICATION	I-squared	PROTECTION FROM BIAS	Bias reason*	OVERALL
APOE	Cau only	e2/3/4	B	616	A	0.000	A		B
COMT	All	rs165599	A	7362	B	0.254	C	Low OR, F	C
COMT	Cau only	rs737865	A	4100	B	0.341	C	Low OR, F	C
DAO	All	rs3918346 (MDAAO-5)	A	1871	A	0.000	C	Low OR	C
DRD1	All	rs4532 (48A/G)	A	1089	A	0.000	A		A
DRD2	Cau only	rs1801028 (Ser311Cys)	B	264	A	0.161	A		B
DRD2	Cau only	rs6277 (Pro319Pro)	A	1276	A	0.153	C	Regr	C
DRD4	All	120-bp TR	A	1050	A	0.071	C	F	C
DRD4	All	rs1800955 (521T/C)	A	3326	A	0.000	C	Low OR	C
DTNBP1	Cau only	rs1011313 (P1325)	A	1002	A	0.000	A		A
GABRB2	Cau only	rs1816071	A	1629	A	0.000	C	Regr	C
GABRB2	Cau only	rs1816072	A	1621	A	0.000	C	Regr	C
GABRB2	Cau only	rs194072	B	603	A	0.074	A		B
GABRB2	Cau only	rs6556547	B	182	A	0.000	A		B
GRIN2B	All	rs1019385 (200T/G)	B	911	B	0.437	C	F	C
GRIN2B	All	rs7301328 (366G/C)	A	1468	A	0.000	C	F	C
HP	All	Hp 1/2	A	2658	A	0.000	C	Low OR	C
IL1B	Cau only	rs16944 (C511T)	A	1533	B	0.256	C	Regr	C
MTHFR	Cau only	rs1801131 (A1298C)	A	1773	A	0.000	A		A
MTHFR	All	rs1801133 (C677T)	A	4999	C	0.555	A		C
PLXNA2	All	rs752016	A	1208	B	0.328	C	F	C
SLC6A4	All	5-HTTVNTR	A	2931	C	0.504	C	F, HWE	C
TP53	All	rs1042522	A	2171	A	0.000	C	Low OR	C
TPH1	All	rs1800532 (218A/C)	A	1937	A	0.129	A		A

# Comparison of SzGene results to previously published meta-analyses

Shaded results indicate discrepancies between prior and SzGene analyses.														
Gene	Study	Published ID	Polymorphism	Prior Meta Sample Size (# Samples)	SzGene Sample Size (# Samples)	Model (default = random effects)	Meta OR	Het Meta	SzGene OR	Het SzGene Result	SzGene Result	HuGE?	Inclusion Date Limit	
APOE	Xu, 2006	16567081	APOE c234/c234	4809 (17)	7995 (24)	c4 v non-c4	1.08 [0.88-1.33]	0.007	1.10 [0.94-1.29]	0.015	N	P	N	Sep-06
APOE	Xu, 2006	16567081	APOE c234/c234	4809 (17)	7995 (24)	c4 v non-c4 (genotypic dominant)	1.09 [0.87-1.36]	n.g.	1.10 [0.92-1.31]	0.019	N	P	N	Sep-06
APOE	Xu, 2006	16567081	APOE c234/c234	5223 (11)	4202 (15)	c4 v non-c4, Caucasian	1.23 [1.04-1.44]	0.59	1.17 [1.01-1.35]	0.615	N	P	N	Sep-06
APOE	Xu, 2006	16567081	APOE c234/c234	5223 (11)	4202 (15)	c4 v non-c4, Caucasian (genotypic dominant)	1.25 [1.04-1.49]	n.g.	1.19 [1.01-1.38]	0.7	N	P	N	Sep-06
APOE	Schuhoff, 2003	12707932	APOE c234/c234	4303 (15)	7995 (24)	**c2 v non-c2	[0.77-1.10]	n.g.	0.99 [0.87-1.13]	0.014	N	P	N	n.g.
APOE	Schuhoff, 2003	12707932	APOE c234/c234	3318 (12)	4202 (15)	**c2 v non-c2, Caucasian	n.g.	n.g.	0.90 [0.75-1.10]	0.67	N	P	N	n.g.
APOE	Schuhoff, 2003	12707932	APOE c234/c234	4303 (15)	7995 (24)	**c3 v non-c3	[0.88-1.11]	n.g.	0.89 [0.83-1.16]	<0.001	N	P	N	n.g.
APOE	Schuhoff, 2003	12707932	APOE c234/c234	3318 (12)	4202 (15)	**c3 v non-c3, Caucasian	n.g.	n.g.	0.92 [0.82-1.04]	0.59	N	P	N	n.g.
APOE	Schuhoff, 2003	12707932	APOE c234/c234	4303 (15)	7995 (24)	**d4 v non-d4	[0.92-1.22]	n.g.	1.18 [1.06-1.31]	0.015	N	P	N	n.g.
APOE	Schuhoff, 2003	12707932	APOE c234/c234	3318 (12)	4202 (15)	**d4 v non-d4, Caucasian	n.g.	n.g.	1.17 [1.01-1.35]	0.615	N	P	N	n.g.
APOE	Schuhoff, 2003	12707932	APOE c234/c234	985 (3)	3211 (5)	**e2 v non-e2, Asian	[0.44-0.91]	n.g.	1.19 [0.80-1.19]	<0.001	N	P	N	n.g.
APOE	Schuhoff, 2003	12707932	APOE c234/c234	985 (3)	3211 (5)	**e3 v non-e3, Asian	1.53 [1.20-1.97]	n.g.	0.90 [0.78-1.03]	<0.001	N	P	N	n.g.
APOE	Schuhoff, 2003	12707932	APOE c234/c234	985 (3)	3211 (5)	**e4 v non-e4, Asian	[0.50-0.95]	n.g.	1.22 [1.02-1.46]	<0.001	N	P	N	n.g.
BDNF	Zinckars, 2007	17413445	270/C/T	1356 (4)	3009 (6)	T v C, Caucasian	1.54 [0.86-2.76]	0.06	1.27 [0.88-1.83]	0.14	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1356 (4)	3009 (6)	(TT + TC) v CC, Caucasian	1.54 [0.85-2.94]	0.06	1.30 [0.88-1.91]	0.13	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1886 (5)	4091 (8)	TT v (TC + CC)	1.30 [0.31-1.57]	0.91	2.11 [0.20-23.65]	n.a.	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1356 (4)	3009 (6)	TT v (TC + CC), Caucasian	1.23 [0.27-5.63]	0.8	2.11 [0.20-23.65]	n.a.	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1886 (5)	4091 (8)	**TT v (TC + CC)	1.43 [0.23-8.60]	0.91	2.11 [0.20-23.65]	n.a.	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1356 (4)	3009 (6)	***TT v (TC + CC), Caucasian	1.43 [0.23-8.60]	0.8	2.11 [0.20-23.65]	n.a.	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1886 (5)	4091 (8)	T v C	1.63 [1.01-2.65]	0.07	1.29 [0.94-1.78]	0.11	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1886 (5)	4091 (8)	**T v C	1.54 [1.14-2.07]	0.07	1.24 [0.94-1.57]	0.11	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1356 (4)	3009 (6)	**T v C, Caucasian	1.43 [1.03-1.99]	0.06	1.21 [0.92-1.69]	0.14	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1886 (5)	4091 (8)	(TT + TC) v CC	1.64 [1.02-2.79]	0.06	1.13 [0.94-1.84]	0.09	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1886 (5)	4091 (8)	**(TT + TC) v CC	1.57 [1.18-2.16]	0.06	1.26 [0.98-1.61]	0.09	P	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	270/C/T	1356 (4)	3009 (6)	**TT v CC, Caucasian	1.44 [1.04-2.06]	0.06	1.23 [0.93-1.63]	0.13	P	N	N	Mar-06
Xu, 2007		17289348	270/C/T	1483 (5)	3009 (6)	C v T, Caucasian	0.74 [0.48-1.14]	0.06	0.79 [0.55-1.14]	0.14	N	N	N	Jul-06
Xu, 2007		17289348	270/C/T	3281 (8)	4091 (8)	C v T	0.59 [0.36-0.96]	<0.001	0.77 [0.56-1.06]	0.11	N	N	N	Jul-06
BDNF	Jonsson, 2006	16581172	270/C/T	2328 (6)	4091 (8)	**T v C	1.35 [1.03-1.77]	n.g.	1.24 [0.98-1.57]	0.11	P	N	N	Oct-05
BDNF	Kanazawa, 2007	17417060	rs6285	6990 (13)	13307 (21)	G v A	1.00 [0.89-1.11]	0.07	1.02 [0.96-1.09]	0.22	N	N	N	Feb-06
BDNF	Zinckars, 2007	17413445	rs6285	3001 (5)	13307 (21)	G v A	1.04 [0.88-1.32]	0.05	1.02 [0.96-1.09]	0.22	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1574 (3)	6287 (9)	G v A, Caucasian	1.24 [0.96-1.60]	0.19	1.05 [0.93-1.20]	0.05	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1427 (2)	6650 (10)	G v A, Asian	0.94 [0.81-1.06]	0.02	1.01 [0.94-1.08]	0.73	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1427 (2)	6650 (10)	**G v A, Asian	0.94 [0.81-1.06]	0.02	1.01 [0.95-1.08]	0.73	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	2518 (3)	12352 (18)	G v A, excl HWE	1.16 [0.88-1.54]	0.02	1.02 [0.95-1.10]	0.14	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	2518 (3)	12352 (18)	**G v A, excl HWE	1.15 [0.98-1.35]	0.02	1.01 [0.96-1.08]	0.14	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	3001 (5)	13307 (21)	(GG + AG) v AA	0.81 [0.68-1.01]	0.07	0.95 [0.85-1.08]	0.38	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1574 (3)	6287 (9)	(GG + AG) v AA, Caucasian	1.00 [0.58-1.83]	0.7	0.83 [0.63-1.09]	0.51	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	3001 (5)	13307 (21)	**GG v AG v AA	0.81 [0.65-1.01]	0.07	0.95 [0.86-1.08]	0.38	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1574 (3)	6287 (9)	**GG v AG v AA, Caucasian	1.04 [0.59-1.83]	0.7	0.83 [0.63-1.09]	0.51	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	2518 (3)	12352 (18)	(GG + AG) v AA, excl HWE	0.84 [0.65-1.08]	0.8	0.95 [0.84-1.07]	0.6	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	2518 (3)	12352 (18)	**GG v AG v AA, excl HWE	0.84 [0.65-1.08]	0.8	0.95 [0.84-1.07]	0.6	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	3001 (5)	13307 (21)	GG v (AG + AA)	1.23 [0.97-1.57]	0.09	1.05 [0.97-1.15]	0.25	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1674 (3)	6287 (9)	GG v (AG + AA), Caucasian	1.20 [0.89-1.66]	0.08	1.09 [0.93-1.27]	0.078	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1427 (2)	6650 (10)	GG v (AG + AA), Asian	1.08 [0.85-1.37]	0.36	1.04 [0.93-1.18]	0.82	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1427 (2)	6650 (10)	**GG v (AG + AA), Asian	1.08 [0.85-1.37]	0.36	1.04 [0.93-1.18]	0.82	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	2518 (3)	12352 (18)	GG v (AG + AA), excl HWE	1.29 [0.97-1.73]	0.06	1.08 [0.96-1.16]	0.19	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	3001 (5)	13307 (21)	**G v A	1.27 [1.06-1.53]	0.05	1.02 [0.96-1.08]	0.22	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1574 (3)	6287 (9)	**G v A, Caucasian	1.27 [1.06-1.53]	0.19	1.03 [0.93-1.14]	0.08	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1427 (2)	6650 (10)	(GG + AG) v AA, Asian	0.77 [0.61-0.96]	0.37	0.98 [0.85-1.14]	0.24	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1427 (2)	6650 (10)	**GG v AG v AA, Asian	0.77 [0.61-0.96]	0.37	0.99 [0.88-1.12]	0.24	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	3001 (5)	13307 (21)	**GG v (AG + AA)	1.23 [1.05-1.44]	0.09	1.05 [0.97-1.13]	0.25	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	1574 (3)	6287 (9)	**GG v (AG + AA), Caucasian	1.37 [1.11-1.69]	0.08	1.07 [0.96-1.20]	0.078	N	N	N	Mar-06
BDNF	Zinckars, 2007	17413445	rs6285	2518 (3)	12352 (18)	**GG v (AG + AA), excl HWE	1.26 [1.07-1.50]	0.06	1.05 [0.97-1.14]	0.19	N	N	N	Mar-06
BDNF	Xu, 2007	17289348	rs6285	7112 (11)	13307 (21)	G v A	0.99 [0.87-1.14]	0.002	1.02 [0.96-1.09]	0.22	N	N	N	Jul-06
BDNF	Xu, 2007	17289348	rs6285	3831 (6)	6287 (9)	G v A, Caucasian	1.03 [0.80-1.33]	0.002	1.05 [0.93-1.20]	0.08	N	N	N	Jul-06
BDNF	Xu, 2007	17289348	rs6285	3281 (5)	6650 (10)	G v A, Asian	0.96 [0.82-1.13]	0.061	1.01 [0.94-1.08]	0.73	N	N	N	Jul-06
BDNF	Zinckars, 2007	17413445	rs6285	1574 (3)	6287 (9)	**GG v (AG + AA), Asian	1.24 [1.05-1.37]	0.07	1.04 [0.96-1.27]	0.27	N	N	N	Mar-06

# Summary

- ① SzGene provides a comprehensive and up-to-date synopsis of genetic association studies in schizophrenia
- ② SzGene almost doubles the number of meta-analyzed genes, highlighting eight novel significant findings
- ③ Genetic effect sizes are generally small (ORs ~1.25)
- ④ Current samples employed in association studies have very little power to detect typical summary ORs
- ⑤ Application of Venice interim guidelines yields four associations with “strong epidemiologic credibility” (*DRD1*, *DTNBP1*, *MTHFR*, *TPH1*)

# **Outlook to part 2 (Session III):**

## **1. Examples of online database structure**

⇒ Example: SzGene (Schizophrenia)

## **2. Inclusion and analysis of GWA studies**

⇒ Example: PDGene (Parkinson's disease)

## **3. Testing “Top Results” in family-based samples**

⇒ Example: AlzGene (Alzheimer's disease)

# Database Content & Development



L. Bertram



K. Mullin



M. McQueen



D. Blacker



N.C. Allen



S. Bagade



R.E. Tanzi

## Online Database Design (Alzheimer Res. Forum)

Colin Knep, Paula Noyes, June Kinoshita

## Online Database Hosting (Schizophrenia Res. Forum)

Alden Bumstead, Hakon Heimer

## Human Genetic Epidemiology Network (HuGENet):

Muin Khoury (National Office for Public Health Genomics, CDC)

John P. Ioannidis (University of Ioannina School of Medicine, Ioannina, Greece)

